Title: ADENO-ASSOCIATED VIRUS VECTORS

## In the Claims

The claims are not amended herein, but are presented below for convenience:

## 1-39. (Cancelled)

- 40. (Previously Presented) A composition comprising:
  - a) a first adeno-associated virus vector comprising linked:
    - i) a first nucleic acid segment comprising a 5'-inverted terminal repeat of adeno-associated virus;
    - a second nucleic acid segment comprising a portion of a gene which includes a transcriptional regulatory region;
    - iii) a third nucleic acid segment comprising a splice donor site; and
    - iv) a fourth nucleic acid segment comprising a 3'-inverted terminal repeat of adeno-associated virus; and
  - b) a second adeno-associated virus vector comprising linked:
    - i) a first nucleic acid segment comprising a 5'-inverted terminal repeat of adeno-associated virus;
    - ii) a second nucleic acid segment comprising a splice acceptor site;
    - a third nucleic acid segment comprising a portion of a gene which together with the nucleic acid segment of a)(ii) comprises a gene comprising an open reading frame which encodes a functional polypeptide; and
    - iv) a fourth nucleic acid segment comprising a 3'-inverted terminal repeat of adeno-associated virus.
- 41. (Previously Presented) The composition of claim 40 further comprising a delivery vehicle.
- 42. (Previously Presented) The composition of claim 40 wherein the nucleic acid segment of a)ii) comprises a promoter.

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- 43. (Previously Presented) The composition of claim 40 wherein the nucleic acid segment of a)ii) comprises an enhancer.
- 44. (Previously Presented) A method to express a polypeptide in a host cell comprising contacting the host cell with the composition of claim 40 so as to express the functional polypeptide.
- 45. (Previously Presented) A method to express a polypeptide in a host cell, comprising: contacting a host cell comprising a first adeno-associated virus vector comprising linked:
  - a) i) a first nucleic acid segment comprising a 5'-inverted terminal repeat of adeno-associated virus;
    - ii) a second nucleic acid segment comprising a portion of a gene which includes a transcriptional regulatory region;
    - iii) a third nucleic acid segment comprising a splice donor site; and
    - iv) a fourth nucleic acid segment comprising a 3'-inverted terminal repeat of adeno-associated virus;

with a second adeno-associated virus vector comprising linked:

- b) i) a first nucleic acid segment comprising a 5'-inverted terminal repeat of adeno-associated virus;
  - ii) a second nucleic acid segment comprising a splice acceptor site;
  - iii) a third nucleic acid segment comprising a portion of a gene which together with the nucleic acid segment of a)ii) comprises a gene comprising an open reading frame which encodes a functional polypeptide; and
  - iv) a fourth nucleic acid segment comprising a 3'-inverted terminal repeat of adeno-associated virus;

so as to yield a host cell which expresses the functional polypeptide.

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46. (Previously Presented) A method to express a polypeptide in a host cell, comprising: contacting a host cell comprising a first adeno-associated virus vector comprising linked:

- a) i) a first nucleic acid segment comprising a 5'-inverted terminal repeat of adeno-associated virus;
  - ii) a second nucleic acid segment comprising a splice acceptor site;
  - iii) a third nucleic acid segment comprising a portion of a gene; and
  - iv) a fourth nucleic acid segment comprising a 3'-inverted terminal repeat of adeno-associated virus;

with a second adeno-associated virus vector comprising linked:

- b) i) a first nucleic acid segment comprising a 5'-inverted terminal repeat of adeno-associated virus;
  - ii) a second nucleic acid segment comprising a portion of a gene which includes a transcriptional regulatory region and which together with the nucleic acid segment of a)iii) comprises a gene comprising an open reading frame which encodes a functional polypeptide;
  - iii) a third nucleic acid segment comprising a splice donor site; and
  - iv) a fourth nucleic acid segment comprising a 3'-inverted terminal repeat of adeno-associated virus;

so as to yield a host cell which expresses the functional polypeptide.

- 47. (Previously Presented) A method to express a polypeptide in a host cell, comprising: contacting a host cell with a first adeno-associated virus vector and a second adeno-associated virus vector, wherein the first adeno-associated virus vector comprises linked:
  - a) i) a first nucleic acid segment comprising a 5'-inverted terminal repeat of adeno-associated virus;
    - ii) a second nucleic acid segment comprising a portion of a gene which includes a transcriptional regulatory region;
    - iii) a third nucleic acid segment comprising a splice donor site; and

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iv) a fourth nucleic acid segment comprising a 3'-inverted terminal repeat of adeno-associated virus;

wherein the second adeno-associated virus vector comprises linked:

- b) i) a first nucleic acid segment comprising a 5'-inverted terminal repeat of adeno-associated virus;
  - ii) a second nucleic acid segment comprising a splice acceptor site;
  - a third nucleic acid segment comprising a portion of a gene which together with the nucleic acid segment of a)ii) comprises a gene comprising an open reading frame which encodes a functional polypeptide; and
  - iv) a fourth nucleic acid segment comprising a 3'-inverted terminal repeat of adeno-associated virus;

so as to yield a host cell which expresses the functional polypeptide.

- 48. (Previously Presented) The method of claim 45 or 47 wherein the nucleic acid segment of a)ii) comprises a promoter.
- 49. (Previously Presented) The method of claim 45 or 47 wherein the nucleic acid segment of a)ii) comprises an enhancer.
- 50. (Previously Presented) The method of claim 46 wherein the nucleic acid segment of b)ii) comprises a promoter.
- 51. (Previously Presented) The method of claim 46 wherein the nucleic acid segment of b)ii) comprises an enhancer.
- 52. (Previously Presented) The method of claim 44, 45, 46 or 47 wherein the host cell is a muscle cell, brain cell, retinal cell, liver cell, lung cell or hematopoietic cell.

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53. (Previously Presented) The method of claim 44, 45, 46 or 47 wherein the polypeptide is the cystic fibrosis transmembrane receptor, polypeptide, β-globin, γ-globin, tyrosine hydroxylase, glucocerebrosidase, arylsulfatase A, factor VIII, dystrophin, or erythropoietin.

- 54. (Previously Presented) The method of claim 44, 45, 46 or 47 wherein the host cell is a mammalian cell.
- 55. (Previously Presented) The method of claim 54 wherein the host cell is a muscle cell, brain cell, retinal cell, liver cell, lung cell or hematopoietic cell.
- 56. (Previously Presented) The method of claim 44, 45, 46 or 47 wherein the host cell is selected from the group consisting of an avian cell, a bovine cell, a swine cell, an equine cell, an ovine cell, a canine cell, a feline cell, an amphibian cell, a reptilian cell and a fish cell.
- 57. (Previously Presented) The method of claim 56 wherein the host cell is a muscle cell, brain cell, retinal cell, liver cell, lung cell or hematopoietic cell.
- 58. (Previously Presented) The method of claim 44, 45, 46, or 47 wherein the vectors comprise DNA.
- 59. (Previously Presented) A composition comprising:a first adeno-associated virus vector comprising linked:
  - a) i) a first nucleic acid segment comprising a 5'-inverted terminal repeat of adeno-associated virus;
    - a second nucleic acid segment comprising a portion of a gene which includes a transcriptional regulatory region;
    - iii) a third nucleic acid segment comprising a splice donor site; and

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iv) a fourth nucleic acid segment comprising a 3'-inverted terminal repeat of adeno-associated virus;

which first vector, in the presence of a second adeno-associated virus vector comprising linked:

- b) i) a first nucleic acid segment comprising a 5'-inverted terminal repeat of adeno-associated virus;
  - ii) a second nucleic acid segment comprising a splice acceptor site;
  - iii) a third nucleic acid segment comprising a portion of a gene which together with the nucleic acid segment of a)ii) comprises a gene comprising an open reading frame which encodes a functional polypeptide;
  - iv) a fourth nucleic acid segment comprising a 3'-inverted terminal repeat of adeno-associated virus;

in a host cell yields a RNA transcript which comprises sequences from the first adenoassociated virus vector linked to sequences from the second adeno-associated virus vector, which sequences are positioned so that the splice donor site is 5' to the splice acceptor site, and which transcript is spliced to a mRNA which encodes the functional protein.

- 60. (Previously Presented) A composition comprising:
  - a first adeno-associated virus vector comprising linked:
    - a) i) a first nucleic acid segment comprising a 5'-inverted terminal repeat of adeno-associated virus;
      - ii) a second nucleic acid segment comprising a splice acceptor site;
      - iii) a third nucleic acid segment comprising a portion of a gene; and
      - iv) a fourth nucleic acid segment comprising a 3'-inverted terminal repeat of adeno-associated virus;

which first vector, in the presence of a second adeno-associated virus vector comprising linked:

b) i) a first nucleic acid segment comprising a 5'-inverted terminal repeat of

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adeno-associated virus;

- ii) a second nucleic acid segment comprising a portion of a gene which together with the nucleic acid segment of a)iii) comprises a gene comprising an open reading frame which encodes a functional polypeptide, wherein the portion of the gene of b)ii) includes a transcriptional regulatory region;
- iii) a third nucleic acid segment comprising a splice donor site;
- iv) a fourth nucleic acid segment comprising a 3'-inverted terminal repeat of adeno-associated virus;

in a host cell yields a RNA transcript which comprises sequences from the first adeno-associated virus vector linked to sequences from the second adeno-associated virus vector, which sequences are positioned so that the splice donor site is 5' to the splice acceptor site, and which transcript is spliced to a mRNA which encodes the functional protein.